

IN THE CLAIMS:

Please amend claims 1-35 as shown below in the detailed listing of all claims which are, or were, in this application:

1. (Currently amended) A method for preparing a sol-gel derived SiO₂ monolith, ~~preferably with a minimum diameter of ≥ 0.5 mm,~~ coating, ~~preferably with a thickness of < 0.5 mm,~~ or particle, ~~preferably with a maximum diameter of ≤ 100 μ m,~~ with having a very fast bioresorption rate, said SiO₂ optionally comprising a specific percentage or percentages of a biologically active agent or agents other than the SiO₂ itself with or without protective agent or agents for said biologically active agent or agents, ~~wherein~~ said method comprising preparing a sol-gel derived SiO₂ ~~is prepared~~ from a sol comprising water, an alkoxide or inorganic silicate and a lower alcohol~~[[, i.e. an alcohol]]~~ with ≤ 4 carbons, using a mineral acid or a base as a catalyst, ~~preferably a mineral acid,~~ ~~and~~ aging said sol ~~is aged~~ and dried ~~characterised~~ in that drying said sol, wherein

a) in the sol the starting

i) pH is from 0.05 to 2.5, ~~preferably 1.5 to 2.5,~~ most ~~preferably 2.0,~~

- ii) molar ratio of water to the alkoxide or inorganic silicate is 0.5 to 2.5, ~~preferably 1.5 to 2.5,~~
 - iii) molar ratio of alcohol to the alkoxide or inorganic silicate is ≥ 0.5 , ~~preferably ≥ 1.0 ;~~ and
- b) either,
- i) the sol is, without induced changes of sol composition,
 - let to gel spontaneously at a temperature of ≤ 25 °C or an elevated temperature of 65 °C to 90 °C, ~~preferably at an elevated temperature of 65 °C to 90 °C,~~ or
 - gelation of the sol is done by forced drying of the sol, or
 - ii) a change or changes of sol composition are induced after sol ageing but before gel formation, said change or changes of sol composition optionally comprising addition of said biologically active agent or agents with or without said protective agent or agents, and
- the ratio t/t_{gel} is ≥ 0.005 , ~~preferably ≥ 0.1 , most preferably ≥ 0.9 ,~~ wherein
- t is the ageing time of the sol, i.e. time from preparation of said sol to the induced changes, and

t_{gel} is the time point where the sol would have turned to a gel without the induced changes; and forced drying of the sol is carried out or initiated within a time of ≤ 30 minutes~~[[~~, preferably ≤ 15 minutes, most preferably ≤ 5 minutes, from said induced change or changes~~]]~~.

2. (Currently amended) A method for adjusting the bioresorption rate of sol-gel derived SiO_2 monolith, ~~preferably with a minimum diameter of ≥ 0.5 mm, coating, preferably with a thickness of ≤ 0.5 mm, or particle, preferably with a maximum diameter of ≤ 100 μm~~ , optionally comprising a specific percentage or percentages of a biologically active agent or agents other than the SiO_2 itself with or without protective agent or agents for said biologically active agent or agents, **characterised** ~~in that~~ wherein

- A) a SiO_2 with a very fast bioresorption rate is obtained according to the method of preparing a SiO_2 of claim 1; and
- B) a SiO_2 with a slower bioresorption rate than the very fast bioresorption rate is obtained by correlating a desired biodegradability of a SiO_2 with changes a), b) and/or c) to said method of preparing a SiO_2 , wherein

a) comprises deviating in the sol any of the starting values:

- i) pH,
- ii) molar ratio of water to the alkoxide or inorganic silicate, and/or
- iii) molar ratio of alcohol to the alkoxide or inorganic silicate;

from the values defined in a) i) - iii) of said method of preparing a SiO_2 ;

b) comprises carrying out induced changes by addition of a component or components, including optional addition of the biologically active agent or agents with or without said protective agent or agents, said changes affecting any of the values i) - iii) of a) of said method of preparing a SiO_2 or a) above if applied by

- i) not carrying out forced drying, or
- ii) carrying out or initiating forced drying of the sol later than defined in b) ii) of said method of preparing a SiO_2 ; and

c) comprises deviating the temperature for letting the sol gel spontaneously from the values defined in b) i) of said method of preparing a SiO_2 ; and
a method for preparing the SiO_2 with said changes to the method correlating with the desired biodegradability is carried out for obtaining the SiO_2 with the desired slower biodegradability.

3. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein an alkoxide, ~~preferably tetraethoxysilane (TEOS),~~ is used ~~for preparing to prepare~~ the sol-gel derived SiO_2 .

4. (Currently amended) The method according to claim 2 ~~characterised in that that~~ wherein an inorganic silicate, ~~preferably sodium or potassium silicate,~~ is used ~~for preparing to prepare~~ the sol-gel derived SiO_2 .

5. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein the lower alcohol is ethanol.

6. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein the induced change is selected from the group consisting of adding water, adding the alkoxide or inorganic silicate, adding the alcohol, adjusting pH by adding an acid or base, ~~preferably the acid or base used as the catalyst,~~ adding the optional bioactive agent or agents with or without protective agent or agents for said biologically active agent or agents affecting any of the values i) - iii) of a) of said method of preparing a SiO₂ or a) of said method of adjusting the bioresorption of sol-gel derived SiO₂ if applied, and any combination thereof.

7. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein ~~drying of~~ the sol is ~~drying~~ dried by a member of the group consisting of ambient heat, vacuum drying, electromagnetic drying, acoustic drying, spray-drying ~~or and~~ freeze-drying, ~~preferably spray-drying or freeze-drying.~~

8. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein the sol is dried by forced drying ~~of~~

~~the sol is carried out, preferably by spray-drying or freeze-drying.~~

9. (Currently amended) The method according to claim 8 ~~characterised in that~~ wherein forced drying is freeze-drying initiated by freezing the sol.

10. (Currently amended) The method according to claim 8 ~~characterised in that~~ wherein the temperature of the sol is $\leq +90\text{ }^{\circ}\text{C}$ [[, preferably $\leq +50\text{ }^{\circ}\text{C}$, most preferably $\leq +40\text{ }^{\circ}\text{C}$]].

11. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein the gel is dried.

12. (Currently amended) The method according to claim 11 ~~characterised in that~~ wherein ~~drying of~~ the gel is ~~drying~~ dried by a member of the group consisting of ambient heat, vacuum drying, electromagnetic drying, acoustic drying, spray-drying ~~or~~ and freeze-drying, ~~preferably ambient heat or freeze-drying.~~

13. (Currently amended) The method according to claim 11 ~~characterised in that~~ wherein the gel is dried at a temperature of $\leq 700\text{ }^{\circ}\text{C}$, ~~preferably $\leq 50\text{ }^{\circ}\text{C}$, and most preferably $\leq 40\text{ }^{\circ}\text{C}$.~~

14. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein a value to be deviated to obtain a slower bioresorption rate is the ratio of water to the alkoxide or inorganic silicate, and the more the ratio of water to alkoxide or inorganic silicate is deviated to be higher or lower the slower the bioresorption rate obtained.

15. (Currently amended) The method according claim 2 ~~characterised in that~~ wherein a value to be deviated to obtain a slower bioresorption rate is the ratio of alcohol to the alkoxide or inorganic silicate, and the more the ratio is deviated to be higher or lower the slower the bioresorption rate obtained.

16. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein a value to be deviated to obtain a slower bioresorption rate is the pH, and the more the pH is

deviated to be higher or lower the slower the bioresorption rate obtained.

17. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein a biologically active agent or agents is added to the sol before gel formation.

18. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein any of the values pH, molar ratio of water to the alkoxide or inorganic silicate, and/or molar ratio of alcohol to the alkoxide or inorganic silicate is changed to deviate from the ranges defined in said method of preparing a SiO_2 , a) ic) - iii), after sol ageing but before gel formation and/or optional addition of said biologically active agent or agents, and within ≤ 30 minutes[[, preferably ≤ 15 minutes and most preferably ≤ 5 minutes]] from the change forced drying of the sol is carried out or initiated.

19. (Currently amended) The method according to claim 2 ~~characterised in that~~ wherein the biologically active agent or agents is selected from the group consisting of a drug, peptide,

protein, hormone, growth factor, enzyme, polysaccharide, living or dead cells or viruses or parts thereof, plasmids, polynucleotides, water soluble ions, salts and any combination thereof.

20. (Currently amended) A bioresorbable sol-gel derived SiO_2 , obtainable according to the method of claim 2, ~~characterised in that~~ wherein

- a) the SiO_2 is a monolith[[, preferably with a minimum diameter of ≥ 0.5 mm]],
- b) the SiO_2 comprises no biologically active agent other than the SiO_2 itself, and
- c) the dissolution rate of the SiO_2 in a TRIS buffer at a temperature of $+37^\circ\text{C}$ and pH 7.4 is ≥ 0.04 wt-%/h[[, preferably ≥ 0.07 wt-%/h and more preferably ≥ 0.15 wt-%/h]].

21. (Currently amended) A bioresorbable sol-gel derived SiO_2 , obtainable according to the method of claim 2, ~~characterised in that~~ wherein

- a) the SiO_2 is a monolith[[, preferably with a minimum diameter of ≥ 0.5 mm]],

- b) the SiO_2 comprises at least one biologically active agent other than the SiO_2 itself, and
- c) the dissolution rate of the SiO_2 in a TRIS buffer at a temperature of +37 °C and pH 7.4 is ≥ 0.35 wt-%/h.

22. (Currently amended) A bioresorbable sol-gel derived SiO_2 , obtainable according to the method of claim 2, ~~characterised in that~~ wherein

- a) the SiO_2 is a coating~~[[~~, preferably with a thickness of < 0.5 mm~~]]~~,
- b) the SiO_2 comprises no biologically active agent other than the SiO_2 itself, and
- c) the dissolution rate of the SiO_2 in TRIS buffer at a temperature of +37 °C and pH 7.4 is ≥ 0.04 wt-%/h~~[[~~, preferably ≥ 0.07 wt-%/h and more preferably ≥ 0.15 wt-%/h~~]]~~.

23. (Currently amended) A bioresorbable sol-gel derived SiO_2 , obtainable according to the method of claim 2, ~~characterised in that~~ wherein

- a) the SiO_2 is a coating~~[[~~, preferably with a thickness of < 0.5 mm~~]]~~,

- b) the SiO_2 comprises at least one biologically active agent other than the SiO_2 itself, and
- c) the dissolution rate of the SiO_2 in TRIS buffer at a temperature of +37 °C and pH 7.4 is ≥ 0.04 wt-%/h[[, preferably ≥ 0.07 wt-%/h and more preferably ≥ 0.15 wt-%/h]].

24. (Currently amended) A bioresorbable sol-gel derived SiO_2 , obtainable according to the method of claim 2 ~~characterised in that~~ wherein

- a) the SiO_2 is a particle[[, preferably with a maximum diameter of $\leq 100 \mu\text{m}$]],
- b) the SiO_2 comprises no biologically active agent other than the SiO_2 itself, and
- c) the dissolution rate of the SiO_2 in TRIS buffer at a temperature of +37 °C and pH 7.4 is ≥ 0.04 wt-%/h[[, preferably ≥ 0.07 wt-%/h and more preferably ≥ 0.15 wt-%/h]].

25. (Currently amended) A bioresorbable sol-gel derived SiO_2 , obtainable according to the method of claim 2 ~~characterised in that~~ wherein

- a) the SiO₂ is a particle[[, preferably with a maximum diameter of $\leq 100 \mu\text{m}$]],
- b) the SiO₂ comprises at least one biologically active agent other than the SiO₂ itself, and
- c) the dissolution rate of the SiO₂ in TRIS buffer at a temperature of +37 °C and pH 7.4 is $\geq 0.5 \text{ wt-\%/h}$.

26. (Currently amended) The SiO₂ according to claim 20, ~~characterised in that~~ wherein the dissolution rate of the SiO₂ is $\geq 0.30 \text{ wt-\%/h}$.

27. (Currently amended) The SiO₂ according to claim 21, ~~characterised in that~~ wherein the dissolution rate of the SiO₂ is $\geq 0.5 \text{ wt-\%/h}$ [[preferably $\geq 1.0 \text{ wt-\%/h}$, more preferably $\geq 2.0 \text{ wt-\%/h}$ and most preferably $\geq 4.0 \text{ wt-\%/h}$]].

28. (Currently amended) A bioresorbable sol-gel derived SiO₂, obtainable according to the method of claim 2, ~~characterised in that~~ wherein

- a) the SiO₂ is a monolith[[, preferably with a minimum diameter of $\geq 0.5 \text{ mm}$]],

- b) the SiO_2 comprises no biologically active agent other than the SiO_2 itself, and
- c) the dissolution rate of the SiO_2 in a TRIS buffer at a temperature of +37 °C and pH 7.4 is from 0.001 to 0.15 wt-%/h[[, preferably from 0.002 to 0.07 wt-%/h, and more preferably from 0.006 to 0.05 wt-%/h]].

29. (Currently amended) A bioresorbable sol-gel derived SiO_2 , obtainable according to the method of claim 2, ~~characterised in that~~ wherein

- a) the SiO_2 is a monolith[[, preferably with a minimum diameter of ≥ 0.5 mm]],
- b) the SiO_2 comprises at least one biologically active agent other than the SiO_2 itself, and
- c) the dissolution rate of the SiO_2 in a TRIS buffer at a temperature of +37 °C and pH 7.4 is from 0.001 to 0.06 wt-%/h [[, preferably from 0.002 to 0.05 wt-%/h, and from 0.006 to 0.025 wt-%/h]].

30. (Currently amended) The SiO_2 according to claim 22 ~~characterised in that~~ wherein the dissolution rate of the SiO_2 in

TRIS buffer at a temperature of +37 °C and pH 7.4 is from 0.001 to 0.15 wt-%/h[[, preferably from 0.002 to 0.07 wt-%/h, and more preferably from 0.006 to 0.05 wt-%/h]].

31. (Currently amended) A bioresorbable sol-gel derived SiO₂, obtainable according to the method of claim 2 ~~characterised in that~~ wherein

- a) the SiO₂ is a particle[[, preferably with a maximum diameter of $\leq 100 \mu\text{m}$]],
- b) the SiO₂ comprises no biologically active agent other than the SiO₂ itself, and
- c) the dissolution rate of the SiO₂ in TRIS buffer at a temperature of +37 °C and pH 7.4 is from 0.001 to 0.008[[, and preferably from 0.002 to 0.003 wt-%/h]].

32. (Currently amended) A bioresorbable sol-gel derived SiO₂, obtainable according to the method of claim 2 ~~characterised in that~~ wherein

- a) the SiO₂ is a particle[[, preferably with a maximum diameter of $\leq 100 \mu\text{m}$]],

- b) the SiO_2 comprises at least one biologically active agent other than the SiO_2 itself, and
- c) the dissolution rate of the SiO_2 in TRIS buffer at a temperature of +37 °C and pH 7.4 is from 0.001 to 0.10 wt-%/h **[[**, preferably from 0.002 to 0.07 wt-%/h, and more preferably from 0.006 to 0.05 wt-%/h**]]**.

33. (Currently amended) A bioresorbable sol-gel derived SiO_2 monolith, ~~preferably with a minimum diameter of ≥ 0.5 mm~~, coating, ~~preferably with a thickness of < 0.5 mm~~, or particle, ~~preferably with a maximum diameter of ≤ 100 μm~~ , obtainable according to the method of claim 2, wherein said SiO_2 comprises a biologically active agent other than the SiO_2 itself and said biologically active agent is a peptide, protein or cell, ~~characterised in that~~ wherein the dissolution rate of the SiO_2 in TRIS buffer at a temperature of +37 °C and pH 7.4 is ≥ 0.04 wt-%/h**[[**, preferably ≥ 0.07 wt-%/h and more preferably ≥ 0.15 wt-%/h**]]**.

34. (Currently amended) A bioresorbable sol-gel derived SiO_2 monolith, ~~preferably with a minimum diameter of ≥ 0.5 mm~~, coating, ~~preferably with a thickness of < 0.5 mm~~, or particle, ~~preferably~~

~~with a maximum diameter of $\leq 100 \mu\text{m}$,~~ obtainable according to the method of claim 2, wherein said SiO_2 comprises a biologically active agent other than the SiO_2 itself and said biologically active agent is a peptide, protein or cell, ~~characterised in that~~ wherein the dissolution rate of the SiO_2 is $\geq 0.5 \text{ wt-\%/h}$ ~~and preferably $\geq 4.0 \text{ wt-\%/h}$.~~

35. (Currently amended) A bioresorbable sol-gel derived SiO_2 monolith, ~~preferably with a minimum diameter of $\geq 0.5 \text{ mm}$,~~ coating, ~~preferably with a thickness of $\leq 0.5 \text{ mm}$,~~ or particle, ~~preferably with a maximum diameter of $\leq 100 \mu\text{m}$,~~ obtainable according to the method of claim 2, wherein said SiO_2 comprises a biologically active agent other than the SiO_2 itself and said biologically active agent is a peptide, protein or cell, ~~characterised in that~~ wherein the dissolution rate of the SiO_2 in TRIS buffer at a temperature of $+37^\circ\text{C}$ and pH 7.4 is from 0.001 to 0.15 wt-%/h[[, preferably from 0.002 to 0.07 wt-%/h, and more preferably from 0.006 to 0.05 wt-%/h]]].

36. (Previously presented) Use of a bioresorbable sol-gel derived SiO_2 according to claim 20 for administering a biologically active

agent to a human or animal body, wherein said use comprises administering selected from the group consisting of oral, buccal, rectal, parenteral, pulmonary, nasal, ocular, intrauterine, vaginal, urethral, topical, transdermal and surgically implantable administering.

37. (Previously presented) Use of a bioresorbable sol-gel derived SiO_2 according to claim 20 for administering a biologically active agent to a plant.